

Package ‘lrgpr’

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Title lrgpr

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Description Fit a Low Rank Gaussian Process Regression (LRGPR) / Linear Mixed Model (LMM) for large datasets. These models are widely used in statistical genetics as a test of association while correcting for the confounding effects of kinship and population structure.

Depends R (>= 3.0.0), methods, Rcpp, RcppGSL, RcppProgress, MASS, parallel, doParallel, formula.tools, BH, bigmemory (>= 4.4.7), biganalytics, aod

LinkingTo Rcpp, RcppGSL

URL <http://lrgpr.r-forge.r-project.org/>

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Collate 'genericFunctions.R' 'lrgpr.R' 'plink.R' 'plots.R'

R topics documented:

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AIC.lrgpr	<i>Akaike’s Information Criterion (AIC)</i>
-----------	---

Description

AIC for model fit by [lrgpr](#)

Usage

`AIC.lrgpr(object, ..., k = 2)`

Arguments

- | | |
|---------------------|--------------------------------------|
| <code>object</code> | model fit with lrgpr |
| <code>...</code> | other arguments |
| <code>k</code> | for compotability, not used |

BIC.lrgpr	<i>Bayesian Information Criterion (BIC)</i>
-----------	---

Description

BIC for model fit by [lrgpr](#)

Usage

```
BIC.lrgpr(object, ...)
```

Arguments

object	model fit with lrgpr
...	other arguments

coefficients.lrgpr	<i>Extract Model Coefficients</i>
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Description

Coefficients estimated with [lrgpr](#)

Usage

```
coefficients.lrgpr(object)
```

Arguments

object	model fit with lrgpr
...	other arguments

convertToBinary	<i>Convert ASCII to binary file</i>
-----------------	-------------------------------------

Description

Converts TPED/DOSAGE/GEN files to binary format

Usage

```
convertToBinary(filename, filenameOut, format,
               nthreads = detectCores(logical = TRUE))
```

Arguments

filename	file to be converted
filenameOut	name of binary file produced
format	specify 'TPED', 'DOSAGE' or 'GEN'
nthreads	number of threads to use

Details

- TPED: plink file can be in either `--recode` or `--recode12` format
- DOSAGE: file follows plink format: <http://pngu.mgh.harvard.edu/~purcell/plink/dosage.shtml>

Example:

```
SNP A1 A2 F1 I1 F2 I2 F3 I3
rs0001 A C 0.98 0.02 1.00 0.00 0.00 0.01
rs0002 G A 0.00 1.00 0.00 0.00 0.99 0.01
```

where the F* values correspond to the dosage values

- GEN: file follows OXFORD format

cooks.distance.lrgpr	<i>Regression Deletion Diagnostics</i>
----------------------	--

Description

Basic quantities for regression deletion diagnostics from fit of `lrgpr`

Usage

```
cooks.distance.lrgpr(model,
  infl = lm.influence(model, do.coef = FALSE),
  res = weighted.residuals(model),
  sd = sqrt(deviance(model)/df.residual(model)),
  hat = infl$hat, ...)
```

Arguments

model	model fit with <code>lrgpr</code>
infl	influence structure as returned by <code>lm.influence</code>
res	residuals
sd	standard deviation to use
hat	hat values
...	other arguments

criterion.lrgpr	<i>Compute AIC/BIC/GCV for <code>lrgpr</code> model as rank changes</i>
-----------------	---

Description

Evaluate information criteria to select an optimal rank for model fit by `lrgpr`

Usage

```
criterion.lrgpr(formula, features, order,
  rank = c(seq(1, 10), seq(20, 100, by = 10), seq(200, 1000, by = 100)))
```

Arguments

formula	standard linear modeling syntax as used in ‘lm’
features	matrix from which the SVD is performed
order	sorted indices of features. When rank is 10, <code>decomp = svd(X[,order[1:10]])</code>
rank	array with elements indicating the number of confounding covariates to be used in the random effect.

See Also

`plot.criterion.lrgpr`, `cv.lrgpr`

Examples

```

n = 300
p = 5000
X = matrix(sample(0:2, n*p, replace=TRUE), nrow=n)

dcmp = svd(X)

# simulate response
h_sq = .8
eta = dcmp$u[,1:2] %*% rgamma(2, 2, 1)
error_var = (1-h_sq) / h_sq * var(eta)
y = eta + rnorm(n, sd=sqrt(error_var))

# Get ordering based on marginal correlation
i = order(cor(y, X)^2, decreasing=TRUE)

# Fit AIC / BIC / GCV based on degrees of freedom
fit = criterion.lrgpr( y ~ 1, features=X, order=i)

plot(fit)

```

cv.lrgpr

Cross-validation for lrgpr

Description

Fit cross-validation for multiple ranks of [lrgpr](#)

Usage

```

cv.lrgpr(formula, features, order, nfolds = 10,
  rank = c(seq(0, 10), seq(20, 100, by = 10), seq(200, 1000, by = 100)),
  nthreads = 1)

```

Arguments

formula	standard linear modeling syntax as used in ‘lm’
features	matrix from which the SVD is performed
order	sorted indices of features. When rank is 10, decomp = svd(X[,order[1:10]])
nfolds	number of training sets
rank	array with elements indicating the number of confounding covariates to be used in the random effect.
nthreads	number of threads to be used

Examples

```

n = 300
p = 5000
X = matrix(sample(0:2, n*p, replace=TRUE), nrow=n)

dcmp = svd(X)

# simulate response
h_sq = .8
eta = dcmp$u[,1:2] %*% rgamma(2, 2, 1)
error_var = (1-h_sq) / h_sq * var(eta)
y = eta + rnorm(n, sd=sqrt(error_var))

# Get ordering based on marginal correlation
i = order(cor(y, X)^2, decreasing=TRUE)

# Fit cross-validation
fit = cv.lrgpr( y ~ 1, features=X, order=i)

plot(fit)

```

df.residual.lrgpr *Residual Degrees-of-Freedom*

Description

Residual df from fit of [lrgpr](#)

Usage

```
df.residual.lrgpr(object, ...)
```

Arguments

object	model fit with lrgpr
...	other arguments

error.bar *Plot Error Bars*

Description

Plot error bars for a confidence interval

Usage

```
error.bar(x, y, upper, lower = upper, length = 0.1, ...)
```

Arguments

x	x-axis position
y	y-axis position
upper	height of bar above y
lower	height of bar below y
length	horizontal length of the error bar
...	arguments for arrows(...)

getAlleleFreq	<i>Calculate allele frequency</i>
---------------	-----------------------------------

Description

Calculate allele frequency

Usage

```
getAlleleFreq(X, nthreads = detectCores(logical = TRUE),
              progress = TRUE)
```

Arguments

X	matrix where each column is a marker coded 0,1,2 or with dosage values in this range
nthreads	number of threads to use
progress	show progress bar

getAlleleVariance	<i>Evaluate variance for each column</i>
-------------------	--

Description

Evaluate variance for each column

Usage

```
getAlleleVariance(X,
                  nthreads = detectCores(logical = TRUE),
                  progress = TRUE)
```

Arguments

X	matrix where each column is a marker
nthreads	number of threads to use
progress	show progress bar

getMACHrsq	<i>Evaluate MACH r^2 information metric for each column</i>
------------	--

Description

Evaluate MACH r^2 information metric for each column

Usage

```
getMACHrsq(X, nthreads = detectCores(logical = TRUE),
           progress = TRUE)
```

Arguments

X	matrix where each column is a marker
nthreads	number of threads to use
progress	show progress bar

See definition in Supplementary information S3 for Marchini and Howie (2010):
<http://www.nature.com/nrg/journal/v11/n7/extref/nrg2796-s3.pdf>.
 #'

References

Marchini, J. and B. Howie. (2010) Genotype imputation for genome-wide association studies. *Nature Reviews Genetics* 11, 499-511 '

getMissingCount	<i>Count missing values</i>
-----------------	-----------------------------

Description

Count missing values

Usage

```
getMissingCount(X,
                nthreads = detectCores(logical = TRUE),
                progress = TRUE)
```

Arguments

X	matrix where each column is a marker
nthreads	number of threads to use
progress	show progress bar

glmApply

*Fit standard linear or logistic model for many markers***Description**

Analogous to [lrgprApply](#), but fits standard linear or logistic models for many markers

Usage

```
glmApply(formula, features, terms = NULL,
         family = gaussian(), useMean = TRUE,
         nthreads = detectCores(logical = TRUE),
         univariateTest = TRUE, multivariateTest = FALSE,
         verbose = FALSE, progress = TRUE, cincl = c(),
         cexcl = c())
```

Arguments

formula	standard linear modeling syntax as used in 'lm'. SNP is a place holder for the each successive column of features
features	a matrix where the statistical model is evaluated with SNP if formula replace by each column successively
terms	indices of the coefficients to be tested. The indices corresponding to SNP are used if terms is not specified
family	gaussian() for a continuous response, and binomial() to fit a logit model for a binary response
useMean	if TRUE, replace missing entries with column mean. Otherwise, do not evaluate the model for that column
nthreads	number of to use for parallel execution
univariateTest	perform univariate hypothesis test for each response for each feature in the loop variable
multivariateTest	perform multivariate hypothesis test for each response (if more than one) for each feature. Note that the runtime is cubic in the number of response variables
verbose	print additional information
progress	show progress bar
cincl	column indeces of features to include for analysis
cexcl	column indeces of features to exclude for analysis

Examples

```

# Generate data
n = 100
p = 500
X = matrix(sample(0:2, n*p, replace=TRUE), nrow=n)
y = rnorm(n)
sex = as.factor(sample(1:2, n, replace=TRUE))

# Fit model for all markers
pValues = glmApply( y ~ sex + sex:SNP, features=X, terms=c(3,4))

# Multivariate model
n = 100
p = 1000
m = 10

Y = matrix(rnorm(n*m), nrow=n, ncol=m)
X = matrix(rnorm(n*p), nrow=n, ncol=p)

res = glmApply( Y ~ SNP, features = X, terms=2, multivariateTest=TRUE)

# p-values for univariate hypothesis test of each feature against
# each response
res$pValues

# p-values for multivariate hypothesis test of each feature against
# all responses are the same time
# returns the results of the Hotelling and Pillai tests
res$pValues_mv

# The multivariate test for X[,1]
res$pValues_mv[1,]

# The result is the same as the standard tests in R
fit = manova( Y ~ X[,1])

summary(fit, test="Hotelling-Lawley")
summary(fit, test="Pillai")

```

glmApply2

Like glmApply, by linear instead of quadratic as a function of the number of covariates

Description

Like glmApply, by linear instead of quadratic as a function of the number of covariates

Usage

```
glmApply2(formula, features, terms = NULL,
  family = gaussian(), useMean = TRUE,
  nthreads = detectCores(logical = TRUE),
  univariateTest = TRUE, multivariateTest = FALSE,
  verbose = FALSE, progress = TRUE, cincl = c(),
  cexcl = c())
```

influence.lrgpr	<i>Regression Diagnostics</i>
-----------------	-------------------------------

Description

Basic quantities for regression diagnostics from fit of `lrgpr`

Usage

```
influence.lrgpr(model, ...)
```

Arguments

model	model fit with <code>lrgpr</code>
...	other arguments

leverage.lrgpr	<i>Regression Diagnostics</i>
----------------	-------------------------------

Description

Basic quantities for regression diagnostics from fit of `lrgpr`

Usage

```
leverage.lrgpr(object)
```

Arguments

object	model fit with <code>lrgpr</code>
--------	-----------------------------------

`lm.influence.lrgpr` *Regression Diagnostics*

Description

Basic quantities for regression diagnostics from fit of `lrgpr`

Usage

```
lm.influence.lrgpr(object, ...)
```

Arguments

<code>object</code>	model fit with <code>lrgpr</code>
<code>...</code>	other arguments

`logLik.lrgpr` *Extract Log-Likelihood*

Description

Log-Likelihood for model fit by `lrgpr`

Usage

```
logLik.lrgpr(object, ...)
```

Arguments

<code>object</code>	model fit with <code>lrgpr</code>
<code>...</code>	other arguments

<code>loss.lrgpr</code>	<i>Loss function</i>
-------------------------	----------------------

Description

Compare observed and fitted response under some loss function

Usage

```
loss.lrgpr(y, yhat, family)
```

Arguments

<code>y</code>	observed response
<code>yhat</code>	fitted response
<code>family</code>	"gaussian" or "binomial"

<code>lrgpr</code>	<i>Fit a Low Rank Gaussian Process Regression (LRGPR) / Linear Mixed Model (LMM)</i>
--------------------	--

Description

Fit LRGPR/LMM models that account for covariance in response values, but where the scale of the covariance is unknown. Standard linear modeling syntax is used for the model specification in addition to a covariance matrix or its eigen-decomposition.

Usage

```
lrgpr(formula, decomp,
      rank = max(ncol(decomp$u), ncol(decomp$vectors)),
      delta = NULL, nthreads = 4, W_til = NULL, scale = TRUE,
      diagnostic = FALSE)
```

Arguments

<code>formula</code>	standard linear modeling syntax as used in ‘lm’
<code>decomp</code>	eigen-decomposition produced from <code>eigen(K)</code> , where <code>K</code> is the covariance matrix. Or singular value decomposition <code>svd(X[,1:100])</code> based on a subset of markers
<code>rank</code>	decomposition is truncated to the first rank eigen-vectors
<code>delta</code>	ratio of variance components governing the fit of the model. This should be estimated from a previous evaluation of ‘lm’ on the same response and eigen-decomposition
<code>nthreads</code>	number of threads to use for parallel execution

W_til	markers used to construct decomp that should now be removed from construction of decomp. This is the proximal contamination term of Listgarten, et al. (2012)
scale	should W_til be scaled and centered
diagnostic	compute diagnostic statistics to be used with plot()

Value

coefficients	regression coefficients for each covariate
p.values	p-values from Wald test of each coefficient
sd	standard deviation of each coefficient estimate
sigSq_e	variance component σ_e^2
	corresponding to the residual error
sigSq_a	variance component σ_a^2
	corresponding the scale of the covariance, K
delta	ratio of variance components: σ_e^2 / σ_a^2
rank	the rank of the random effect
logLik	log-likelihood of the model fit
fitted.values	estimated response values: y_hat
alpha	BLUP of the random effect
Sigma	variance-covariate matrix of estimate of beta
hii	diagonals of the matrix H such that y_hat = Hy
y	responses
x	design matrix
df	effective degrees of freedom: trace(H) based on Hoffman (2013)
residuals	residuals of model fit: y - y_hat
AIC	Akaike information criterion
BIC	Bayesian information criterion
GCV	generalized cross-validation
eigenVectors	eigen-vectors in decomp
eigenValues	eigen-values in decomp
df.residual	n - ncol(X)
rank	rank of decomposition used, where only non-negative eigen/singular values are considered

Details

`lrgpr` fits the model:

$$y = X\beta + \alpha + \epsilon$$

$$\alpha \sim N(0, K\sigma_a^2)$$

$$\epsilon \sim N(0, \sigma_e^2)$$

where

$$\delta = \sigma_e^2 / \sigma_a^2$$

In practice the eigen-decomposition of K , and not K itself is required. The rank can be set to use only eigen-vectors 1:rank in the model.

This package allows hypothesis tests of single coefficients using `fit$p.values` which fits a Wald test. Composite hypothesis tests of multiple coefficients are performed with `wald(fit, terms=1:3)`.

Note that likelihood ratio tests with linear mixed models do not perform well and the resulting p-values often do not follow a uniform distribution under the null (Pinheiro and Bates, 2000). We strongly advise against using it with this model.

`lrgpr` uses the algorithm of Lippert, et al. (2011).

See Hoffman (2013) for an interpretation of the linear mixed model.

References

- Kang, H. M., et al. (2010) Variance component model to account for sample structure in genome-wide association studies. *Nature Genetics* 42, 348-54
- Lippert, C., et al. (2011) FaST linear mixed models for genome-wide association studies. *Nature Methods* 9, 525-26
- Listgarten, J., et al. (2012) Improved linear mixed models for genome-wide association studies. *Nature Methods* 8, 833-5
- Rasmussen, C. E. and Williams, C. K. I. (2006) Gaussian processes for machine learning. MIT Press
- Pinheiro, J. C. and Bates, D. M. (2000) Mixed-Effects Models in S and S-PLUS. Springer, New York
- Hoffman, G. E. (2013) Correcting for Population Structure and Kinship Using the Linear Mixed Model: Theory and Extensions. *PLoS ONE* 8(10):e75707
- Note that degrees freedom and some diagnostic statistics are not currently calculated when `W_til` is specified.

See Also

`wald`, `lrgprApply`

Examples

```
# Generate random data
set.seed(1)
n <- 200
y <- rnorm(n)
K <- crossprod( matrix(rnorm(n*1000), ncol=n) )
age <- rpois(n, 50)
sex <- as.factor(sample(1:2, n, replace=TRUE))
decomp <- eigen(K)

# Fit the model
fit <- lrgpr( y ~ sex + age, decomp, diagnostic=TRUE)

# Print results
fit

# Print more detailed results
summary(fit)

# P-values for each covariate
fit$p.values

# Visualize fit of the model like for `lm'
par(mfrow=c(2,2))
plot(fit)

# Composite hypothesis test using Wald's test
# Joint test of coefficients 2:3
wald( fit, terms=2:3)
```

lrgprApply

Fit a Low Rank Gaussian Process Regression (LRGPR) / Linear Mixed Model (LMM) for many markers

Description

Fit LRGPR/LMM models that account for covariance in response values, but where the scale of the covariance is unknown. It returns p-values equivalent to the results of `lrgpr` and `lrgpr`, but is designed to analyze thousands of markers in a single function call.

Usage

```
lrgprApply(formula, features, decomp, terms = NULL,
  rank = max(ncol(decomp$u), ncol(decomp$vectors)),
  map = NULL, distance = NULL, dcmp_features = NULL,
  W_til = NULL, scale = TRUE, delta = NULL,
  reEstimateDelta = FALSE,
  nthreads = detectCores(logical = TRUE),
  verbose = FALSE, progress = TRUE, cincl = c(),
  cexcl = c())
```

Arguments

formula	standard linear modeling syntax as used in 'lm'. SNP is a place holder for the each successive column of features
features	a matrix where the statistical model is evaluated with SNP if formula replace by each column successively
decomp	eigen-decomposition produced from eigen(K), where K is the covariance matrix. Or singular value decomposition svd(features[,1:100]) based on a subset of markers
terms	indices of the coefficients to be tested. The indices corresponding to SNP are used if terms is not specified
rank	decomposition is truncated to the first rank eigen-vectors
map	p x 2 matrix where each entry corresponds to a marker in features. First column is the marker names, second columns is the genetic or physical location
distance	size of the proximal contamination window in units specified by map.
dcmp_features	the indices in features of the markers used to construct dcmp
W_til	markers used to construct decomp that should now be removed from construction of decomp. This is the proximal contamination term of Listgarten, et al. (2012)
scale	should W_til be scaled and centered
delta	ratio of variance components governing the fit of the model. This should be estimated from a previous evaluation of 'lm' on the same response and eigen-decomposition
reEstimateDelta	should delta be re-estimated for every marker. Note: reEstimateDelta=TRUE is much slower
nthreads	number of to use for parallel execution
verbose	print extra information
progress	show progress bar
cincl	column indices of features to include for analysis
cexcl	column indices of features to exclude for analysis

Examples

```
# Generate data
n = 100
p = 500
X = matrix(sample(0:2, n*p, replace=TRUE), nrow=n)
y = rnorm(n)
sex = as.factor(sample(1:2, n, replace=TRUE))

K = tcrossprod(matrix(rnorm(n*n*3), nrow=n))
decomp = eigen(K, symmetric=TRUE)

# Fit null model
```

```
fit = lrgpr( y ~ sex, decomp)

# Fit model for all markers
pValues = lrgprApply( y ~ sex + sex:SNP, features=X, decomp, terms=c(3,4), delta=fit$delta)
```

```
plot.criterion.lrgpr
```

Plot AIC/BIC/GCV values for lrgpr() model as rank changes

Description

Plots the criteria metrics returned by `criterion.lrgpr`

Usage

```
plot.criterion.lrgpr(x, col = rainbow(3), ...)
```

Arguments

<code>x</code>	list returned by <code>criterion.lrgpr</code>
<code>col</code>	array of 3 colors
<code>...</code>	other arguments

See Also

`criterion.lrgpr`

```
plot.cv.lrgpr
```

Plot Results of Cross-validation

Description

Plot results of `cv.lrgpr`, which fits cross-validation for multiple ranks of the LRGPR

Usage

```
plot.cv.lrgpr(x,
  ylim = c(min(x$cvse - x$cvse), max(x$cvse + x$cvse)),
  xlim = range(x$rank), pch = 20, col = "red",
  main = "Cross validation", xlab = "# of markers used",
  ylab = "Cross validation error", ...)
```

Arguments

x	result of cv.lrgpr
ylim	limits of y-axis
xlim	limits of x-axis
pch	pch
col	col
main	main
xlab	xlab
ylab	ylab
...	other parameters fed to plot()

plot.lrgpr

*Plot Diagnostics for an lrgpr Object***Description**

Six plots (selectable by "which") are currently available: a plot of residuals against fitted values, a Scale-Location plot of $\sqrt{| \text{residuals} |}$ against fitted values, a Normal Q-Q plot, a plot of Cook's distances versus row labels, a plot of residuals against leverages, and a plot of Cook's distances against leverage/(1-leverage). By default, the first three and "5" are provided.

Usage

```
plot.lrgpr(x, which = c(1L:3L, 5L),
  caption = list("Residuals vs Fitted", "Normal Q-Q", "Scale-Location", "Cook's d"),
  panel = if (add.smooth) panel.smooth else points,
  sub.caption = NULL, main = "",
  ask = prod(par("mfcol")) < length(which) && dev.interactive(),
  ..., id.n = 3, labels.id = names(residuals(x)),
  cex.id = 0.75, qqline = TRUE, cook.levels = c(0.5, 1),
  add.smooth = getOption("add.smooth"),
  label.pos = c(4, 2), cex.caption = 1)
```

Arguments

x	lrgpr object.
which	if a subset of the plots is required, specify a subset of the numbers "1:6".
caption	captions to appear above the plots; "character" vector or "list" of valid graphics annotations, see "as.graphicsAnnot". Can be set to "" or "NA" to suppress all captions.
panel	panel function. The useful alternative to "points", "panel.smooth" can be chosen by "add.smooth = TRUE".

sub.caption	common title-above the figures if there are more than one; used as "sub" (s."title") otherwise. If "NULL", as by default, a possible abbreviated version of "deparse(x\$call)" is used.
main	title to each plot-in addition to "caption".
ask	logical; if "TRUE", the user is <code>_ask_ed</code> before each plot, see "par(ask=.)".
...	other parameters to be passed through to plotting functions.
id.n	number of points to be labelled in each plot, starting with the most extreme.
labels.id	vector of labels, from which the labels for extreme points will be chosen. "NULL" uses observation numbers.
cex.id	magnification of point labels.
qqline	logical indicating if a "qqline()" should be added to the normal Q-Q plot.
cook.levels	levels of Cook's distance at which to draw contours.
add.smooth	logical indicating if a smoother should be added to most plots; see also "panel" above.
label.pos	positioning of labels, for the left half and right half of the graph respectively, for plots 1-3.
cex.caption	controls the size of "caption".

See Also

`plot.lm`

<code>predict.lrgpr</code>	<i>Predict response</i>
----------------------------	-------------------------

Description

Predict response values after training with `lrgpr`. Leaving `X_test` and `K_test` as `NULL` returns the fitted values on the training set

Usage

```
predict.lrgpr(object, X_test = NULL, K_test = NULL, ...)
```

Arguments

<code>object</code>	model fit from <code>lrgpr</code> on training samples
<code>X_test</code>	design matrix of covariates for test samples
<code>K_test</code>	covariance matrix between samples in the test set and training set
...	other arguments

<code>print.lrgpr</code>	<i>Print Values</i>
--------------------------	---------------------

Description

Print details for fit from `lrgpr`

Usage

```
print.lrgpr(x, ...)
```

Arguments

<code>x</code>	model fit from <code>lrgpr</code>
<code>...</code>	other arguments

<code>print.summary.lrgpr</code>	<i>Object Summaries</i>
----------------------------------	-------------------------

Description

Print summary for fit from `lrgpr`

Usage

```
print.summary.lrgpr(x, ...)
```

Arguments

<code>x</code>	model fit from <code>lrgpr</code>
<code>...</code>	other arguments

QQ_plot

QQ plot

Description

QQ plot and lambda_GC optimized for large datasets.

Usage

```
QQ_plot(p_values,
        col = rainbow(min(length(p_values), ncol(p_values))),
        main = "", pch = 20, errors = TRUE, lambda = TRUE,
        p_thresh = 1e-06, showNames = FALSE, ylim = NULL,
        xlim = NULL, plot = TRUE, new = TRUE,
        box.lty = par("lty"), collapse = FALSE, ...)
```

Arguments

p_values	vector, matrix or list of p-values
col	colors corresponding to the number of columns in matrix, or entries in the list
main	title
pch	pch
errors	show 95% confidence interval
lambda	calculate and show genomic control lambda. Lambda_GC is calculated using the 'median' method on p-values > p_thresh.
p_thresh	Lambda_GC is calculated using the 'median' method on p-values > p_thresh.
showNames	show column names or list keys in the legend
ylim	ylim
xlim	xlim
plot	make a plot. If FALSE, returns lambda_GC values without making plot
new	make a new plot. If FALSE, overlays QQ over current plot
box.lty	box line type
collapse	combine entries in matrix or list into a single vector
...	other arguments

Examples

```
p = runif(1e6)
QQ_plot(p)

# get lambda_GC values without making plot
lambda = QQ_plot(p, plot=FALSE)
```

<code>read.fam</code>	<i>Read plink FAM/TFAM files</i>
-----------------------	----------------------------------

Description

Read FAM/TFAM file into a dataframe. This function is the same as `read.tfam`

Usage

```
read.fam(file)
```

Arguments

<code>file</code>	location of FAM/TFAM file
-------------------	---------------------------

<code>read.tfam</code>	<i>Read plink FAM/TFAM files</i>
------------------------	----------------------------------

Description

Read FAM/TFAM file into a dataframe. This function is the same as `read.fam`

Usage

```
read.tfam(file)
```

Arguments

<code>file</code>	location of FAM/TFAM file
-------------------	---------------------------

<code>residuals.lrgpr</code>	<i>Extract Model Residuals</i>
------------------------------	--------------------------------

Description

Residuals fitted with `lrgpr`

Usage

```
residuals.lrgpr(object, type = "working", ...)
```

Arguments

<code>object</code>	model fit with <code>lrgpr</code>
<code>type</code>	the type of residual, but there is only one option here
<code>...</code>	other arguments

rstandard.lrgpr	<i>Regression Deletion Diagnostics</i>
-----------------	--

Description

Basic quantities for regression deletion diagnostics from fit of `lrgpr`

Usage

```
rstandard.lrgpr(model, ...)
```

Arguments

model	model fit with <code>lrgpr</code>
...	other arguments

set_missing_to_mean	<i>Replace Missing Values with Mean</i>
---------------------	---

Description

For each column, replace NA values with the column mean

Usage

```
set_missing_to_mean(A)
```

Arguments

A	matrix
---	--------

summary.lrgpr	<i>Summarizing LRGPR / Linear Mixed Model Fits</i>
---------------	--

Description

Print summary for fit from `lrgpr`

Usage

```
summary.lrgpr(object, ...)
```

Arguments

object	model fit from <code>lrgpr</code>
...	other arguments

<code>vcov.lrgpr</code>	<i>Calculate Variance-Covariance Matrix for a <code>lrgpr</code> Object</i>
-------------------------	---

Description

Returns the variance-covariance matrix of the main parameters of a fitted model object

Usage

```
vcov.lrgpr(object, ...)
```

Arguments

<code>object</code>	model fit with <code>lrgpr</code>
<code>...</code>	other arguments

<code>wald</code>	<i>Composite hypothesis test of multiple coefficients</i>
-------------------	---

Description

Performs a multi-dimensional Wald test against $H_0: \beta_i \dots \beta_j = 0$ using the estimated coefficients and their variance-covariance matrix

Usage

```
wald(fit, terms)
```

Arguments

<code>fit</code>	result of fitting with <code>lrgpr</code>
<code>terms</code>	indices of the coefficients to be tested

Details

The Wald statistic is

$$\beta_h^T \Sigma_h^{-1} \beta_h \sim \chi^2_{|h|}$$

where

$$h$$

specifies the coefficients being tested and

$$|h|$$

is the number of entries

See Also

`lrgpr`

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